

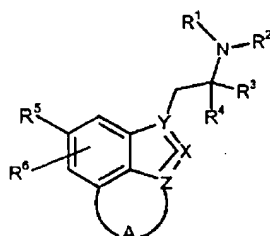
U.S. Patent Application No. 10/721,204
 Amendment dated March 3, 2005
 Response to Office Action dated December 8, 2004

AMENDMENTS TO THE CLAIMS:

This listing of claims will replace all prior versions, and listings, of claims in the application:

LISTING OF CLAIMS:

1. (currently amended) A compound represented by Formula I:



wherein R^1 and R^2 are independently chosen from hydrogen or an alkyl group;

R^3 and R^4 are independently chosen from hydrogen, an alkyl group or R^3 , R^4 and the carbon atom to which they are attached form a cycloalkyl ring, or R^3 and R^4 together represent $(CH_2)_m$ to form a saturated heterocycle;

R^5 is chosen from hydroxyl, alkoxy, alkyl, halogen, or $OC(=O)W$;

R^6 is chosen from hydrogen, halogen, a substituted or unsubstituted alkyl group;

R^7 and R^8 are hydrogen or an alkyl group;

W is a substituted or unsubstituted alkyl group, NR^7R^8 , $N(R^7)CH_2(CH_2)_nN(R^7)(R^8)$, O-alkyl, or a substituted or unsubstituted alkenyl;

m is 3 or 4;

n is 2 or 3;

A is a 6-membered ring containing 6 carbon atoms a 5- to 7-membered ring optionally containing one heteroatom chosen from NR^7 , O, or S;

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~~X is either N or C; Y is N; Z is C;~~

~~Y and Z are either N or C, wherein Y and Z are different; and~~

~~the dashed bonds denote a suitably appointed single and double bond;~~

~~or pharmaceutically acceptable salts or solvates thereof.~~

2. (currently amended) The compound of claim 1, wherein R^1 and R^2 are independently chosen from hydrogen or C_{1-4} alkyl;

R^3 and R^4 are independently chosen from hydrogen, C_{1-4} alkyl or R^3 , R^4 and the carbon atom to which they are attached form a cyclopropyl ring, ~~or R^2 and R^3 together represent $(CH_2)_m$ to form a saturated heterocycle;~~

R^5 is chosen from hydroxyl, C_{1-4} alkoxy, C_{1-4} alkyl, halogen, or $OC(=O)W$;

R^6 is chosen from hydrogen, halogen, C_{1-4} alkyl, C_{1-4} alkyl substituted with halogen;

R^7 and R^8 are hydrogen or C_{1-4} alkyl;

W is C_{1-6} alkyl, NR^7R^8 , $N(R^7)CH_2(CH_2)_nN(R^7)(R^8)$, OC_{1-6} alkyl, C_{1-6} alkyl optionally substituted with halogen, hydroxyl, CO_2C_{1-4} alkyl, $CON(C_{1-4}alkyl)_2$, $C(=NH)NH_2$, $NHC(=NH)NH_2$, or NH_2 , C_{2-4} alkenyl optionally substituted by phenyl, unsubstituted or substituted with one or more of C_{1-4} alkyl, C_{1-4} alkoxy or halogen;

m is 3 or 4;

n is 2 or 3;

A is ~~a 6-membered ring containing 6 carbon atoms~~ a 5- to 7-membered ring optionally containing one heteroatom chosen from NR^7 , O, or S;

~~X is either N or C; Y is N; Z is C;~~

~~Y and Z are either N or C, wherein Y and Z are different; and~~

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the dashed bonds denote a suitably appointed single and double bond;

or pharmaceutically acceptable salts or solvates thereof.

3. (currently amended) The compound of claim 1, wherein ~~said R² and R³ form a saturated~~
(CH₂)_m ~~heterocycle or~~ said R³ and R⁴ together form a cycloalkyl ring.

4. (currently amended) The compound of claim 1, wherein R¹, R², and R³ are hydrogen;
~~or R² and R³ together represent (CH₂)_m to form a pyrrolidine;~~

R⁴ is C₁₋₄alkyl;

R⁵ is chosen from hydroxyl, C₁₋₄alkoxy, or OC(=O)W;

R⁶ is chosen from hydrogen, halogen, C₁₋₄alkyl, C₁₋₄alkyl substituted with halogen;

R⁷ and R⁸ are hydrogen or C₁₋₄alkyl;

W is C₁₋₆alkyl, NR⁷R⁸, C₁₋₆alkyl optionally substituted with halogen, hydroxyl, or
CO₂C₁₋₄alkyl;

m is 3;

A is a 6-membered ring containing 6 carbon atoms ~~optionally containing one heteroatom chosen
from NR⁷ or O;~~

X is ~~either N or C;~~

Y is N and Z is C; and

the dashed bonds denote a suitably appointed single and double bond.

5. (currently amended) The compound of claim 1, wherein the compound is:

2-(2-Aminopropyl)-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-ol;

2-(2-Dimethylaminoethyl)-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-ol;

2-(2-Aminopropyl)-5-methyl-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-ol;

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2-(2-Aminopropyl)-5-fluoro-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-ol;
2-(6-Fluoro-7-methoxy-4,5-dihydro-3*H*-benzo[*cd*]indazol-1-yl)-1-methylethylamine;
Cyclopropanecarboxylic acid 2-(2-aminopropyl)-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-yl ester;
~~1-(2-Aminopropyl)-1,3,4,5-tetrahydro-benzo[*cd*]indol-7-ol;~~
~~1-(2-Aminopropyl)-5*H*-pyrano[4,3,2-*cd*]indazol-7-ol; or~~
~~1-(2-Aminopropyl)-4-methyl-1,3,4,5-tetrahydro-pyrazolo[4,3,2-*de*]isoquinolin-7-ol~~ ———— or
combinations thereof.

6-7. (canceled)

8. (original): A method of controlling normal or elevated intraocular pressure comprising administering a pharmaceutically effective amount of a composition comprising at least one compound of claim 1.

9. (canceled)

10. (original): The method of claim 8, wherein said R³ and R⁴ together form a cycloalkyl ring.

11. (currently amended) The method of claim 8, wherein said compound is 2-(2-Aminopropyl)-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-ol;

2-(2-Dimethylaminoethyl)-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-ol;

2-(2-Aminopropyl)-5-methyl-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-ol;

2-(2-Aminopropyl)-5-fluoro-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-ol;

2-(6-Fluoro-7-methoxy-4,5-dihydro-3*H*-benzo[*cd*]indazol-1-yl)-1-methylethylamine;

Cyclopropanecarboxylic acid 2-(2-aminopropyl)-2,6,7,8-tetrahydro-benzo[*cd*]indazol-4-yl ester;

~~1-(2-Aminopropyl)-1,3,4,5-tetrahydro-benzo[*cd*]indol-7-ol;~~

~~1-(2-Aminopropyl)-5*H*-pyrano[4,3,2-*cd*]indazol-7-ol; or~~

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~~1-(2-Aminopropyl)-4-methyl-1,3,4,5-tetrahydro-pyrazolo[4,3,2-de]isoquinolin-7-ol;~~ or
combinations thereof.

12. (currently amended) The method of claim 8, ~~wherein~~ wherein R^1 , R^2 , and R^3 are hydrogen;
~~or R^2 and R^3 together represent $(CH_2)_m$ to form a pyrrolidine;~~

R^4 is C_{1-4} alkyl;

R^5 is chosen from hydroxyl, C_{1-4} alkoxy, or $OC(=O)W$;

R^6 is chosen from hydrogen, halogen, C_{1-4} alkyl, C_{1-4} alkyl substituted with halogen;

R^7 and R^8 are hydrogen or C_{1-4} alkyl;

W is C_{1-6} alkyl, NR^7R^8 , C_{1-6} alkyl optionally substituted with halogen, hydroxyl, or

CO_2C_{1-4} alkyl;

m is 3;

A is a 6-membered ring optionally containing one heteroatom chosen from NR^7 or O;

X is ~~either~~ N or C;

Y is N and Z is C; and

the dashed bonds denote a suitably appointed single and double bond.

13-14. (canceled)

15. (original): A method for the treatment of glaucoma comprising administering a
pharmaceutically effective amount of a composition comprising at least one compound of claim 1.

16. (currently amended) The method of claim 15, wherein R^1 , R^2 , and R^3 are hydrogen;

~~or R^2 and R^3 together represent $(CH_2)_m$ to form a pyrrolidine;~~

R^4 is C_{1-4} alkyl;

R^5 is chosen from hydroxyl, C_{1-4} alkoxy, or $OC(=O)W$;

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R⁶ is chosen from hydrogen, halogen, C₁₋₄alkyl, C₁₋₄alkyl substituted with halogen;

R⁷ and R⁸ are hydrogen or C₁₋₄alkyl;

W is C₁₋₆alkyl, NR⁷R⁸, C₁₋₆alkyl optionally substituted with halogen, hydroxyl, or
CO₂C₁₋₄alkyl;

m is 3;

A is a 6-membered ring containing 6 carbon atoms ~~optionally containing one heteroatom chosen
from NR⁷ or O;~~

X is ~~either N or C;~~

Y is N and Z is C; and

the dashed bonds denote a suitably appointed single and double bond.

17. (currently amended) The method of claim 15, wherein said compound is:

~~1-(2-Aminopropyl)-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-ol;~~

~~1-((S)-2-Aminopropyl)-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-ol;~~

~~(R)-1-((S)-2-Aminopropyl)-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-ol;~~

~~(S)-1-((S)-2-Aminopropyl)-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-ol;~~

~~1-((S)-2-Aminopropyl)-3-methyl-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-ol;~~

~~1-(S)-1-Pyrrolidin-2-ylmethyl-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-ol;~~

~~1-((S)-2-Aminopropyl)-5-fluoro-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-ol;~~

~~[1-((S)-2-Aminopropyl)-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-yl]-dimethylamine;~~

~~[1-((S)-2-Aminopropyl)-1,7,8,9-tetrahydro-pyrano[2,3-g]indazol-8-yl]-methanol;~~

~~1-(2-Aminopropyl)-1,7,8,9-tetrahydro-pyrano[3,2-g]indazol-8-ol;~~

~~1-(Pyrrolidin-2-ylmethyl)-3,7,8,9-tetrahydro-pyrano[3,2-c]indazol-8-ol;~~

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~~1-((S)-2-Aminopropyl)-3,7,8,9-tetrahydro-pyrano[3,2-e]indazol-8-ol; or~~
~~1-((S)-2-Aminopropyl)-3-methyl-3,7,8,9-tetrahydro-pyrano[3,2-e]indazol-8-ol; 2-(2-~~
Aminopropyl)-2,6,7,8-tetrahydro-benzo[cd]indazol-4-ol;
2-(2-Dimethylaminoethyl)-2,6,7,8-tetrahydro-benzo[cd]indazol-4-ol;
2-(2-Aminopropyl)-5-methyl-2,6,7,8-tetrahydro-benzo[cd]indazol-4-ol;
2-(2-Aminopropyl)-5-fluoro-2,6,7,8-tetrahydro-benzo[cd]indazol-4-ol;
2-(6-Fluoro-7-methoxy-4,5-dihydro-3H-benzo[cd]indazol-1-yl)-1-methylethylamine;
Cyclopropanecarboxylic acid 2-(2-aminopropyl)-2,6,7,8-tetrahydro-benzo[cd]indazol-4-yl ester;
or mixtures thereof.

18. (original): A pharmaceutical composition comprising the compound of claim 1 and at least one carrier.
19. (previously presented) A method to activate serotonin receptors comprising administering an effective amount of at least one compound of claim 1 to a patient.